

## AMENDMENTS

Please amend the claim set to read as follows.

1-42. (canceled)

43. (currently amended) A method for obtaining an siRNA molecule for a target gene, wherein said siRNA molecule comprises an antisense sequence that is 19 – 30 nucleotide bases in length and a sense sequence that is 19 – 30 nucleotide bases in length and said antisense sequence and said sense sequence form a duplex region of 19-30 base pairs, said method comprising the steps:

- (a) selecting a target gene;
- (b) identifying a set of candidate siRNA sequences, wherein the antisense sequence of each of said candidate siRNA sequences is at least 79% complementary to a region of the target gene;
- (c) applying to each of said candidate siRNA sequences a computer algorithm, wherein said computer algorithm comprises a set of one or more criteria selected from the group consisting of a presence of U at position 1 of the antisense sequence or a presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence, a presence of U at position 17 of the antisense sequence or a presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, a presence of A at position 10 of the antisense sequence or a presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence, a presence of U at position 6 of the antisense sequence or a presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence, an absence of G at position 1 of the antisense sequence or an absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, an absence of C at position 7 of the antisense sequence or an absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, an absence of A at position 15 of the antisense sequence or an absence of U at a position of the sense sequence that is complementary

to position 15 of the antisense sequence and an absence of U at position 9 of the antisense sequence or an absence of A at a position of the sense sequence that is complementary to position 9 of the antisense sequence, wherein said positions are defined in reference to the 5' end of the antisense sequence within the duplex region;

- (d) after step (c) selecting a candidate siRNA sequence from the set of candidate siRNA sequences of step (b) as an siRNA sequence for the target gene, wherein if said candidate siRNA sequence satisfies said set of one or more criteria; and
- (e) after step (d) synthesizing an siRNA molecule for said target gene, wherein said siRNA molecule for said target gene comprises said siRNA sequence for the target gene, whereby said siRNA molecule for said target gene is obtained.

44. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence.

45. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence.

46. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the presence of A at position 10 of the antisense sequence or the presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence.

47. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the presence of U at position 6 of the antisense sequence or the

presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence.

48. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence.
49. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence.
50. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the absence of A at position 15 of the antisense sequence or the absence of U at a position of the sense sequence that is complementary to position 15 of the antisense sequence.
51. (currently amended) The method according to claim 43, wherein the set of one or more criteria includes the absence of U at position 9 of the antisense sequence or the absence of A at a position of the sense sequence that is complementary to position 9 of the antisense sequence.
52. (currently amended) The method according to claim 43 further comprising applying one or more additional criteria selected from the group consisting of: a GC content between about 30% and 52%, and at least 2 A or U bases at positions 1 – 5 of the antisense sequence or at least 2 A or U bases at positions of the sense sequence that are complementary to positions 1-5 of the antisense sequence, and selecting said candidate siRNA sequence if said candidate siRNA sequence satisfies said one or more additional criteria and wherein said candidate siRNA that is selected satisfies said one or more additional criteria.

53. (currently amended) The method according to claim 43, wherein said candidate siRNA sequence that is selected as said siRNA sequence for the target gene if said candidate siRNA sequence satisfies at least two criteria selected from the group consisting of: the presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, the presence of A at position 10 of the antisense sequence or the presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence, the presence of U at position 6 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence, the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, the absence of A at position 15 of the antisense sequence or the absence of U at a position of the sense sequence that is complementary to position 15 of the antisense sequence, and the absence of U at position 9 of the antisense sequence or the absence of A at a position of the sense sequence that is complementary to position 9 of the antisense sequence.

54. (currently amended) The method according to claim 43, wherein said candidate siRNA sequence that is selected as said siRNA sequence for the target gene if said candidate siRNA sequence satisfies at least three criteria selected from the group consisting of: the presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, the presence of A at position 10 of the

antisense sequence or the presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence, the presence of U at position 6 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence, the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, the absence of A at position 15 of the antisense sequence or the absence of U at a position of the sense sequence that is complementary to position 15 of the antisense sequence, and the absence of U at position 9 of the antisense sequence or the absence of A at a position of the sense sequence that is complementary to position 9 of the antisense sequence.

55. (canceled)

56. (canceled)

57. (currently amended) The method according to claim 43, wherein in (c) said method comprises applying the following criteria to each of said candidate siRNA sequences, the presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, and further comprises applying each of the following additional criteria to each of the candidate siRNA sequences: a GC content between about 30% and 52%, and at least 2 A or U bases at

positions 1 – 5 of the antisense sequence or at least 2 A or U bases at positions 15-19 of the sense sequence that are complementary to positions 1-5 of the antisense sequence, and in (d) selecting said candidate siRNA sequence as said siRNA sequence that is selected for said target gene if said candidate siRNA sequence satisfies the criteria of the presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, the GC content between about 30% and 52%, and at least 2 A or U bases at position 1 – 5 of the antisense sequence or at least 2 A or U bases at positions 15-19 of the sense sequence that are complementary to positions 1-5 of the antisense sequence.

58. (currently amended) The method according to claim 43, wherein in (c) said method comprises applying the criteria of the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence and further comprises applying the criteria of a GC content of between 30% and 52% and in (d) selecting said candidate siRNA sequence that is selected as said siRNA sequence for said target gene if said siRNA sequence satisfies both of said criteria of: (i) the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence; and (ii) the GC content of between 30% and 52%.

59. (currently amended) The method according to claim 43, wherein said candidate siRNA sequence that is selected as said siRNA sequence for the target gene if said candidate siRNA sequence satisfies each of the following criteria: the absence of G at

position 1 of the antisense sequence and the absence of C at position 7 of the antisense sequence.

60. (currently amended) The method according to claim 43, wherein in (c), said method comprises applying the criteria of the absence of C at position 7 or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence and further comprises applying the criteria of: a GC content of between 30% and 52%, and wherein ~~in (d) said method comprises selecting~~ said candidate siRNA sequence that is selected as said siRNA sequence for said target gene ~~if said siRNA sequence~~ satisfies all of the criteria of: (i) the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence; and the GC content of between 30% and 52%.

61-67 (canceled)

68. (currently amended) A method for selecting an siRNA sequence for a target gene, wherein said siRNA comprises an antisense sequence that is 19 – 30 nucleotide bases in length and a sense sequence that is 19 – 30 nucleotide bases in length and said antisense sequence and said sense sequence form a duplex region of 19-30 base pairs, said method comprising the steps:

- a. selecting a target gene;
- b. identifying a set of candidate siRNA sequences, wherein the antisense sequence of each of said candidate siRNA sequences is at least 79% complementary to a region of the target gene;
- c. accessing a computer and causing said computer to apply ~~applying~~ to each of said candidate siRNA sequences, a computer algorithm, wherein said computer algorithm is stored in computer readable form and comprises a set of one or more criteria selected from the group consisting of a presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense

sequence, a presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, a presence of A at position 10 of the antisense sequence or the presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence, a presence of U at position 6 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence, an absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, an absence of C at position 7 of the antisense sequence or the absence presence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, an absence of A at position 15 of the antisense sequence or the absence of U at a position of the sense sequence that is complementary to position 15 of the antisense sequence and an absence of U at position 9 of the antisense sequence or the absence of A at a position of the sense sequence that is complementary to position 9 of the antisense sequence, wherein said positions are defined in reference to the 5' end of the antisense sequence within the duplex region;

- d. after step (c) selecting a candidate siRNA sequence from the set of candidate siRNA sequences of step (b) as said siRNA sequence for the target gene, wherein if-said candidate siRNA sequence satisfies said set of one or more criteria, and wherein said selecting is performed by said computer; and
- e. after step (d) generating an output comprising said siRNA sequence for the target gene, wherein said generating is performed by said computer and said output is displayed in a form that is readable by a human-computer.

69. (canceled)

70. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the presence of U at position 1 of the antisense sequence or the

presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence.

71. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence.
72. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the presence of A at position 10 of the antisense sequence or the presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence.
73. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the presence of U at position 6 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence.
74. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence.
75. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence.
76. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the absence of A at position 15 of the antisense sequence or the

absence of U at a position of the sense sequence that is complementary to position 15 of the antisense sequence.

77. (currently amended) The method according to claim 68, wherein the set of one or more criteria includes the absence of U at position 9 of the antisense sequence or the absence of A at a position of the sense sequence that is complementary to position 9 of the antisense sequence.

78. (canceled)

79. (previously presented) The method according to claim 43, wherein in (b) said antisense sequence is 100% complementary to said region of said target gene.

80. (canceled)

81. (previously presented) The method according to claim 68, wherein in (b) said antisense sequence is 100% complementary to said region of said target gene.

82. (canceled) .

83. (canceled)

84. (previously presented) The method according to claim 43, wherein said synthesizing comprises chemical synthesis.

85. (previously presented) The method according to claim 43, wherein said synthesizing comprises enzymatic synthesis.

86. (currently amended) A method for obtaining an siRNA molecule for a target gene, wherein said siRNA molecule comprises an antisense sequence that is 19 – 30 nucleotide bases in length and a sense sequence that is 19 – 30 nucleotide bases in

length and said antisense sequence and said sense sequence form a duplex region of 19-30 base pairs, said method comprising the steps:

- a. selecting a target gene;
- b. identifying a set of candidate siRNA sequences, wherein the antisense sequence of each of said candidate siRNA sequences is at least 79% complementary to a region of the target gene;
- c. applying to each of said candidate siRNA sequences a computer algorithm, wherein said computer algorithm comprises a set of four or more criteria selected from the group consisting of: the presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, the presence of A at position 10 of the antisense sequence or the presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence, the presence of U at position 6 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence, the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, the absence of A at position 15 of the antisense sequence or the absence of U at a position of the sense sequence that is complementary to position 15 of the antisense sequence, the absence of U at position 9 of the antisense sequence or the absence of A at a position of the sense sequence that is complementary to position 9 of the antisense sequence, a GC content between about 30% and 52%, and at least 2 A or U bases at positions 1-5 of the antisense sequence or at least 2 A or U bases at positions 15-19 of the sense sequence that are complementary to positions 1-5

of the antisense sequence, wherein said positions are defined in reference to the 5' end of the antisense sequence within said duplex region;

- d. after step (c) selecting a candidate siRNA sequence from the set of candidate siRNA sequences of step (b) as an siRNA sequence for the target gene, wherein if said candidate siRNA sequence satisfies said set of four or more criteria; and
- e. after step (d) synthesizing said siRNA molecule for said target gene, wherein said siRNA molecule for said target gene comprises said siRNA sequence for the target gene, whereby said siRNA molecule for said target gene is obtained.

87. (currently amended) The method according to claim 86, wherein in (c) said method comprises applying a set of five or more criteria selected from the group consisting of: the presence of U at position 1 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the presence of U at position 17 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 17 of the antisense sequence, the presence of A at position 10 of the antisense sequence or the presence of U at a position of the sense sequence that is complementary to position 10 of the antisense sequence, the presence of U at position 6 of the antisense sequence or the presence of A at a position of the sense sequence that is complementary to position 6 of the antisense sequence, the absence of G at position 1 of the antisense sequence or the absence of C at a position of the sense sequence that is complementary to position 1 of the antisense sequence, the absence of C at position 7 of the antisense sequence or the absence of G at a position of the sense sequence that is complementary to position 7 of the antisense sequence, the absence of A at position 15 of the antisense sequence or the absence of U at a position of the sense sequence that is complementary to position 15 of the antisense sequence, the absence of U at position 9 of the antisense sequence or the absence of U at a position of the sense sequence that is complementary to position 9 of the antisense sequence, a GC content between about 30% and 52%, and at least 2 A or U bases at positions 1 - 5 of

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the antisense sequence or at least 2 A or U bases at positions 15-19 of the sense sequence that are complementary to positions 1-5 of the antisense sequence, and wherein in (d) selecting said candidate siRNA sequence that is selected as said siRNA sequence for said target gene ~~if said candidate siRNA sequence~~ satisfies said set of five or more criteria.

88-91 (canceled)